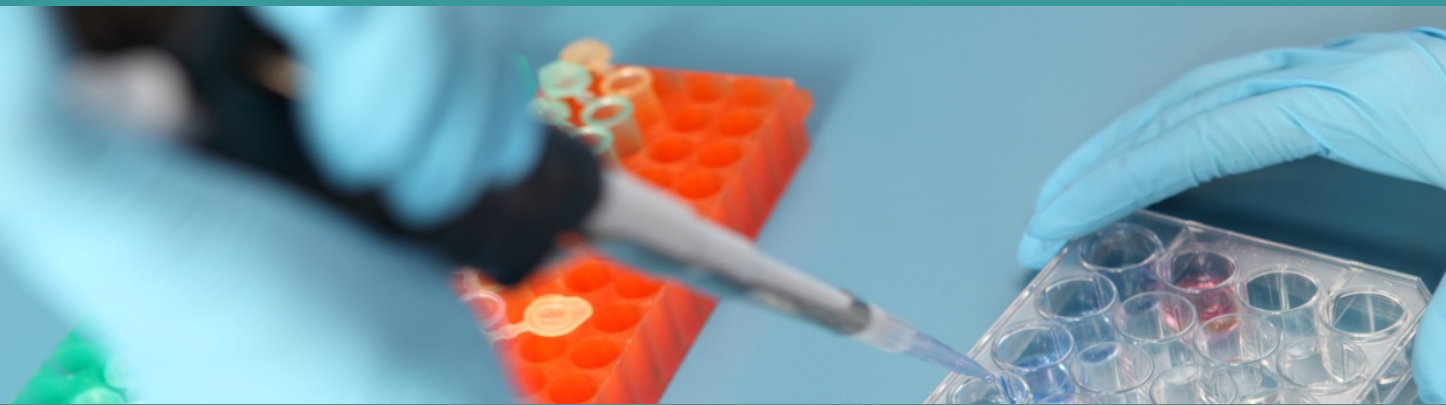




A NEW FRONTIER IN IMMUNO-ONCOLOGY

**LSX Investival Showcase
London, 13th November 2018**

LSE: SCLP.L



A NEW FRONTIER IN IMMUNO-ONCOLOGY



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DIFFERENTIATED IMMUNO-ONCOLOGY CLINICAL STAGE OPPORTUNITY

COMPANY FOCUS

- Scancell is developing innovative immunotherapies for the treatment of cancer

MARKET OPPORTUNITY

- Immuno-oncology is one of the fastest growing sectors in the biopharmaceutical industry (est. CAGR ~20% over next 5 years)

PROPRIETARY TECHNOLOGY PLATFORMS

- Novel immunogenic antigens and modulation mechanisms that stimulate potent T-cell responses for the treatment or prevention of cancer
- Unique mode of action of **IMMUNOBODY**[®] and **MODITOPE**[®] immunotherapies stimulate immune responses by presenting cancer antigens to trigger potent killer T-cell activation

CLINICAL STAGE ASSETS

- Four lead products in development
- Phase II and Phase I/II studies in preparation targeting multiple cancer indications

COMPANY FACTS & FINANCIALS

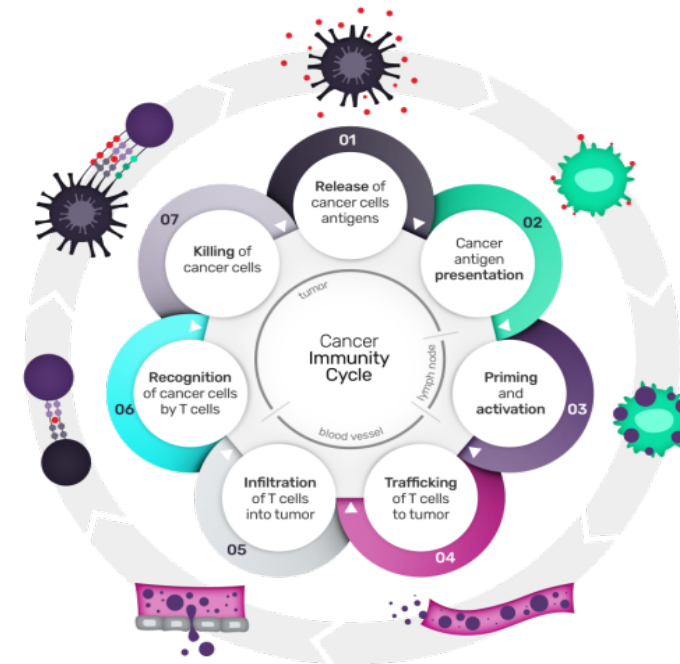
- Scientific founder Professor Lindy Durrant
- Corporate offices based in Oxford, UK
- 23 employees (10 PhD's)
- AIM listed (SCLP)

2 PLATFORMS, 4 LEAD PRODUCTS + MULTIPLE CANCER INDICATIONS



MEETING THE NEED FOR EFFECTIVE THERAPEUTIC CANCER VACCINES

- ▶ Key challenge is to stimulate an effective T cell response to reject or kill the growing tumour
- ▶ Most vaccine strategies only stimulate low frequency, low avidity T cell responses that fail to control tumour growth
- ▶ Scancell's novel therapies stimulate **high avidity** CD8 and/or CD4 T-cells that efficiently kill tumours



Ref: Chen and Mellman 2013

TWO DIFFERENTIATED PLATFORMS

IMMUNOBODY®

- ▶ DNA-based platform generates high avidity CD8 T-cells by presenting T-cell epitopes of known cancer antigens through a unique dual mode of action

MODITOPE®

- ▶ Modified peptides that generate potent killer CD4 T-cells to target antigens induced by stress-induced post-translational modifications (siPTM vaccines)



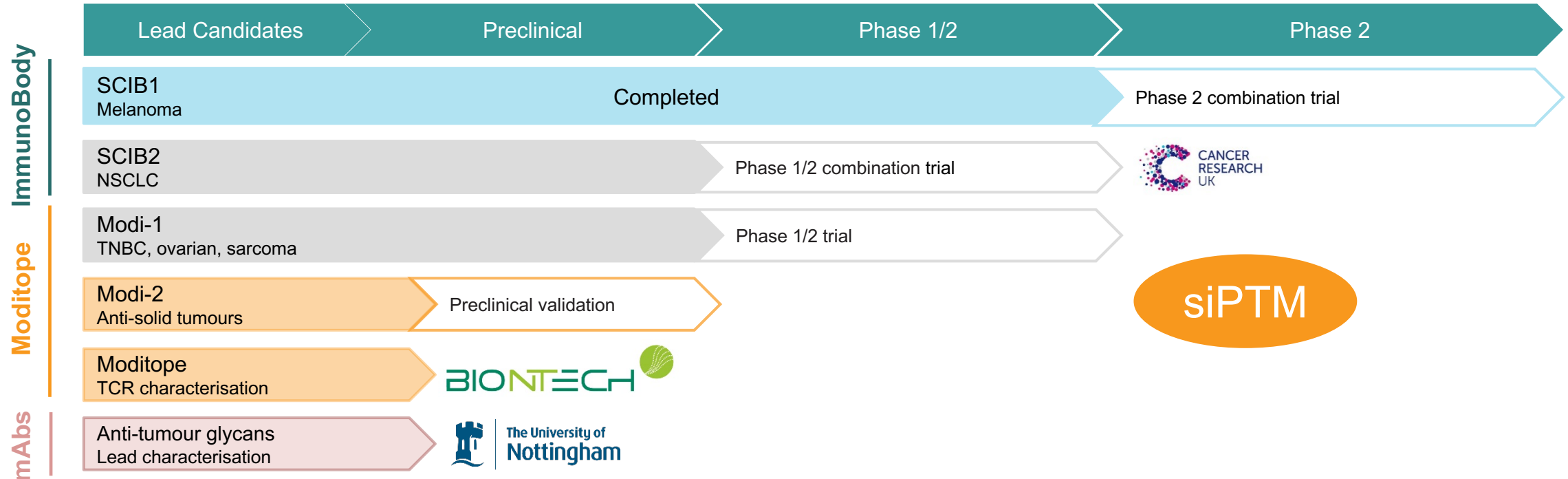
DEVELOPMENT PIPELINE

IMMUNOBODY®

- ▶ **SCIB1:** Targets malignant melanoma. Phase 1/2 study completed with strong survival data. Phase 2 combination trial with immune checkpoint inhibitor planned for 1H CY19
- ▶ **SCIB2:** Targets NSCLC. Phase 1/2 combination trial with immune checkpoint inhibitor to be funded and sponsored by Cancer Research UK (CRUK).

MODITOPE®

- ▶ **Modi-1:** Manufacturing process development initiated. Phase 1/2 trial in TNBC, ovarian and sarcoma planned for 2019.
- ▶ **Modi-2:** Targets multiple solid tumours. Preclinical development of selected epitopes.
- ▶ **TCR collaboration:** To clone and characterise T cell receptors against Modi-1 specific epitopes.



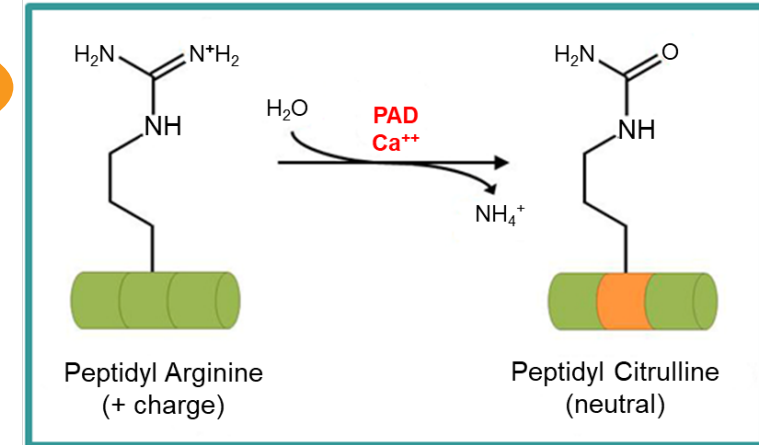


Stress-Induced Post-translational Modifications (siPTM)

- ▶ One such modification involves the process of **CITRULLINATION**
 - ▶ The alteration of proteins due to enzymatic conversion of arginine residues to citrulline
 - ▶ Citrullination occurs as a result of a degradation and 'recycling' process called **autophagy** that is induced in stressed cells, including cancer cells
 - ▶ Citrullinated epitopes presented on **MHC class II**
 - ▶ Patent awarded in Europe, Japan, China, Australia; still being pursued in US but attorney confident we will get broad claims

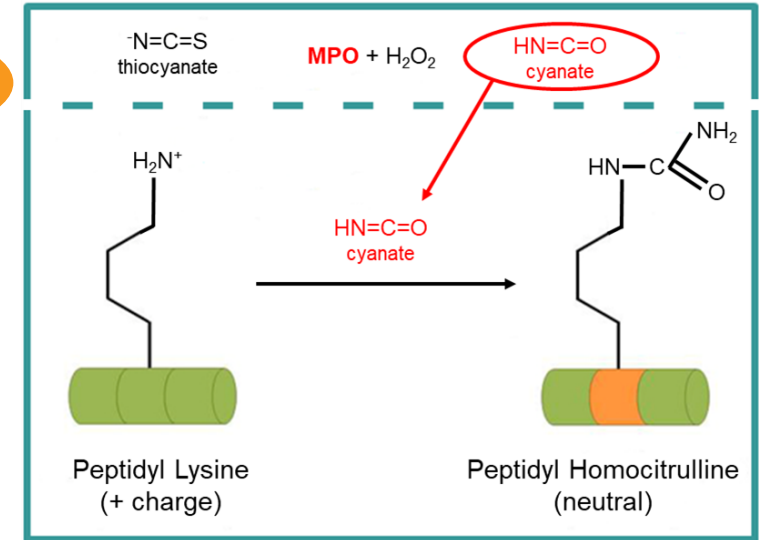
- ▶ Another modification involves the process of **HOMOCITRULLINATION**
 - ▶ The alteration of proteins due to conversion of lysine residues to homocitrulline
 - ▶ Homocitrullination occurs as a result of MPO released by myeloid-derived suppressor cells (MDSC) which converts thiocyanate to cyanate in the presence of H_2O_2
 - ▶ Cyanate diffuses into tumour cells and results in spontaneous homocitrullination of cytoplasmic proteins
 - ▶ These proteins are degraded during **autophagy** and homocitrullinated epitopes presented on **MHC class II**
 - ▶ Patent filed with broad claims in cancer and composition of matter for any use of homocitrullinated peptides

Modi-1



PAD = peptidylarginine deiminase

Modi-2

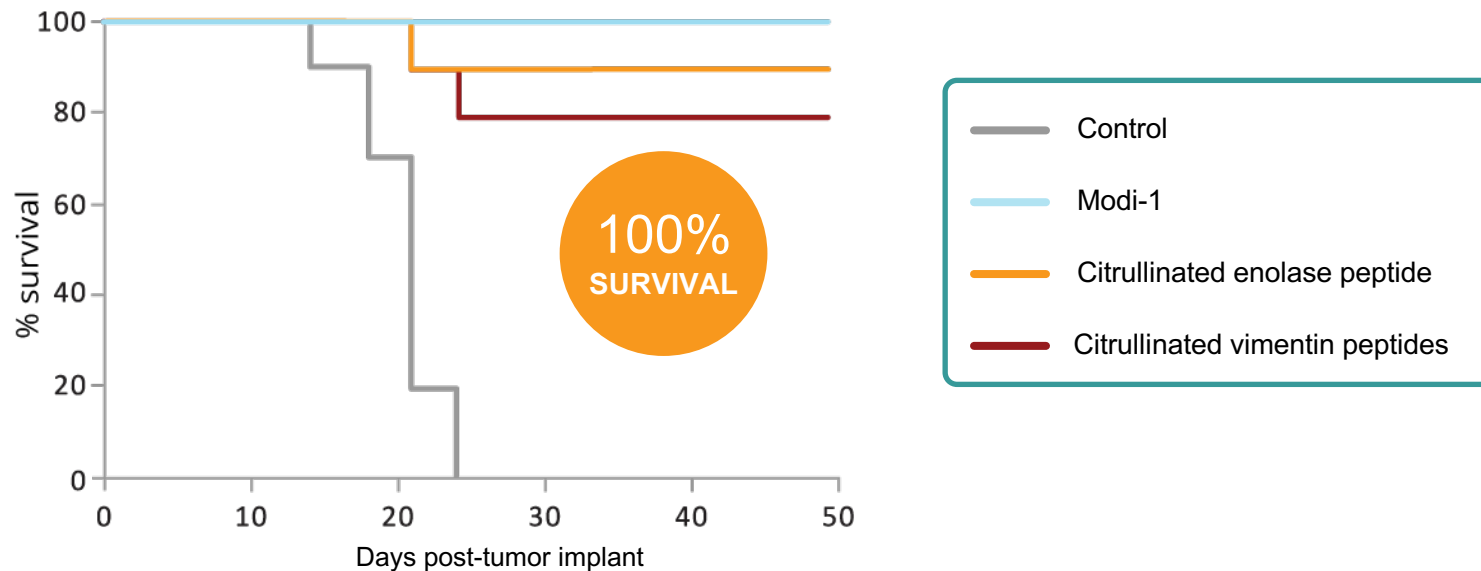


MPO = myeloperoxidase



Modi-1

- ▶ Consists of:
 - ▶ Two citrullinated vimentin peptides (Vim-1 and Vim-2)
 - ▶ One citrullinated enolase peptide (Eno-1)
- ▶ Vimentin and enolase targets are highly expressed in triple negative breast cancer (TNBC) (90%), ovarian cancer (95%), and sarcoma (100%) - all with high unmet medical need
- ▶ Modi-1 induced potent anti-tumour responses in mice with established melanoma (B16)
- ▶ **A single immunization of Modi-1 resulted in a 100% survival rate in animal models**

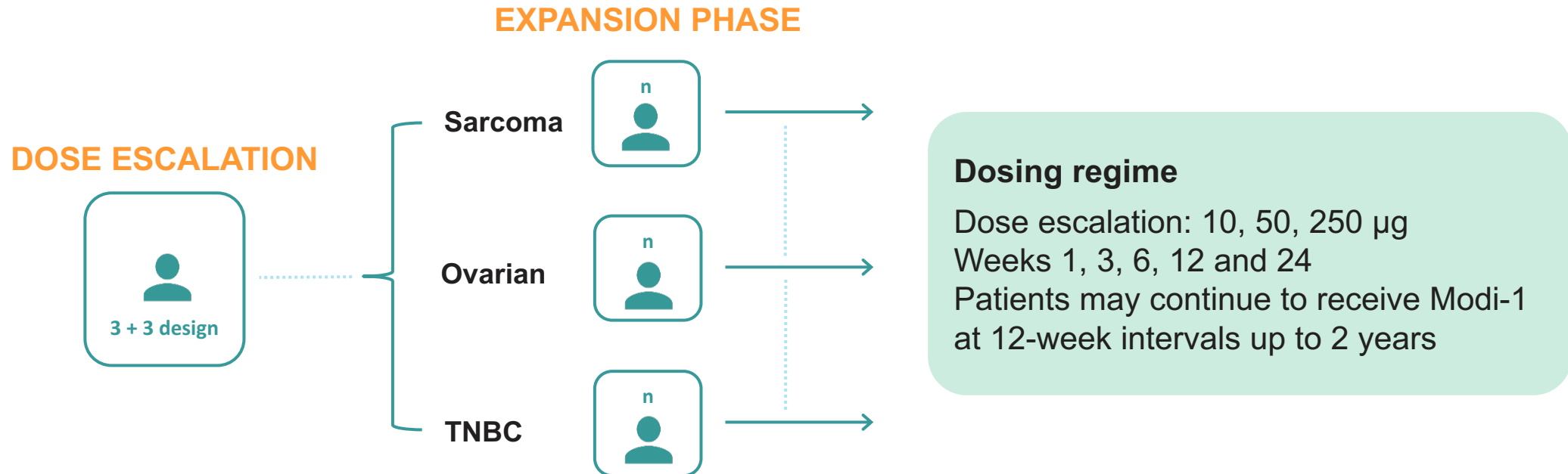




Modi-1 FIRST IN HUMAN STUDY

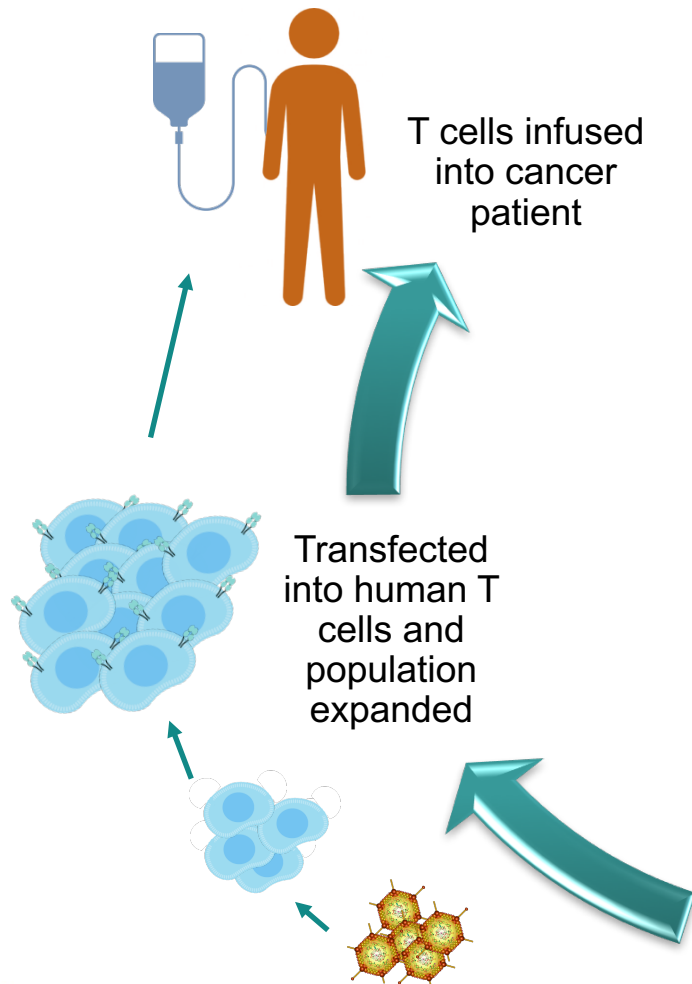
PATIENT POPULATION

- ▶ Patients with tumours with high vimentin or enolase expression (e.g., sarcoma, triple negative breast cancer, ovarian)
- ▶ Failed or intolerant to standard of care therapies





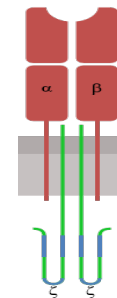
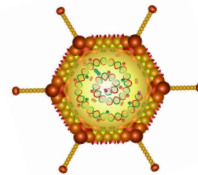
MODITOPE® TCR RESEARCH COLLABORATION



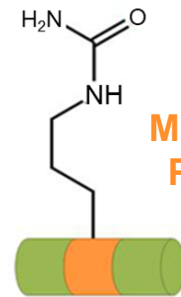
ADVANTAGES OF CITRULLINATED & HOMOCITRULLINATED ANTIGEN-SPECIFIC TCRS

- ▶ Citrullinated & homocitrullinated antigens are expressed by a wide range of tumours
- ▶ Citrullinated & homocitrullinated antigen-specific T cells recognise the non-polymorphic HLA-DP4 so are applicable to at least 70% of patients
- ▶ Citrullinated and homocitrullinated antigen-specific T cells stimulate potent anti-tumour immunity

MODI-1/2
TCR VECTOR



MODI-1/2
TCR





MODITOPE MILESTONES

EXTERNAL VALIDATION OF MODITOPE® PLATFORM INTERNAL PROJECTS ADVANCED AND EXPANDED

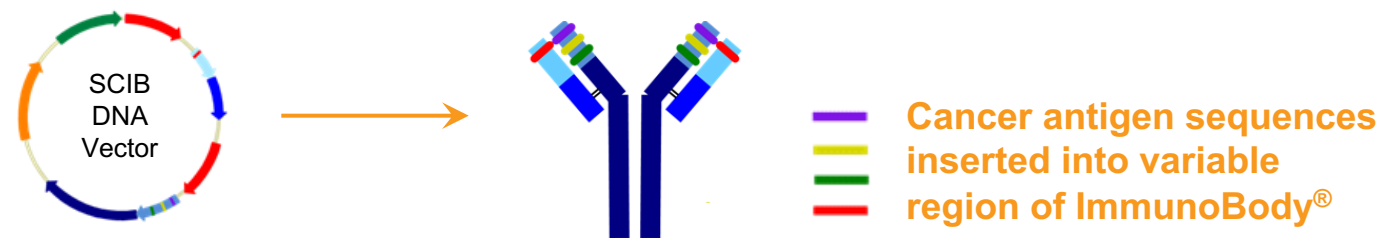
MODITOPE®

- ▶ Research collaboration to develop T-cell based therapies established with BioNTech
- ▶ License agreed with ISA Pharmaceuticals for development of Amplivant® Modi-1 conjugate therapy
- ▶ GMP manufacturer contracted for production of Modi-1/Amplivant® conjugate
- ▶ Modi-1 clinical study expected to start in CY19
- ▶ Homocitrullinated peptides under evaluation for inclusion in new Modi-2 vaccine targeting multiple solid tumours
- ▶ Strong Patent protection
- ▶ Shortlisted for CRUK Grand Challenge award; [Project Blueprint](#)



THE IMMUNOBODY® PLATFORM

- ▶ Proprietary patent protected platform
- ▶ Several cancer associated T cell epitopes are engineered into a human antibody framework to make a genetic antigen/antibody complex
- ▶ Delivered as a DNA plasmid using electroporation



- ▶ Nano-vesicle delivery under evaluation
- ▶ Novel dual mechanism of action based on **direct** and **cross-presentation**
- ▶ SCIB1 for melanoma (**TRP-2/gp100 melanoma associated antigens**): Phase 1/2 clinical trial complete, Phase 2 planned
- ▶ SCIB2 for lung cancer (**NY-ESO-1**): Clinical development partnership with CRUK

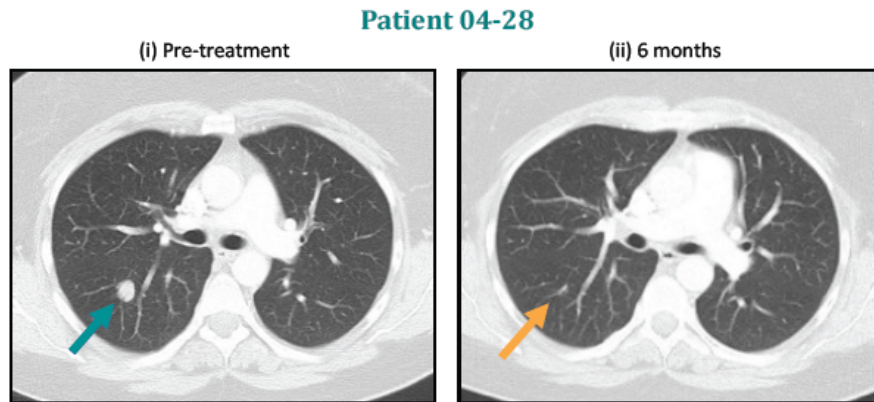


SCIB1 IN PATIENTS WITH MELANOMA

SCIB1 has an excellent safety profile with no dose-limiting toxicities and no serious adverse events related to study drug or delivery device

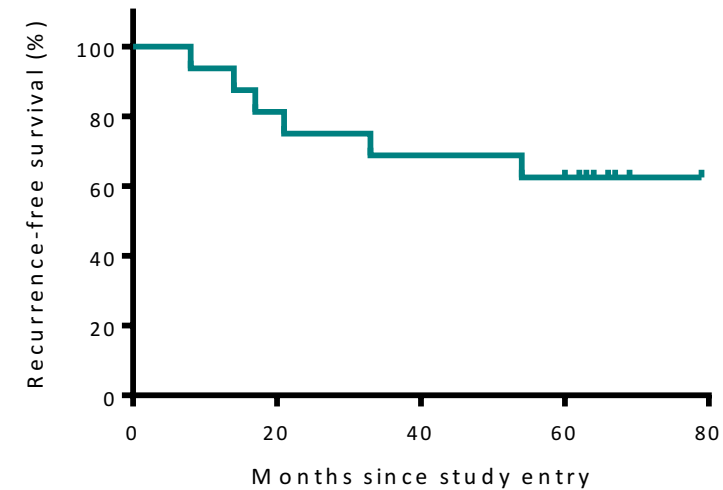
TUMOUR RESPONSE

Patient with tumour received 8 mg SCIB1 and showed a marked reduction in size of detectable lung lesions



SURVIVAL IN RESECTED PATIENTS

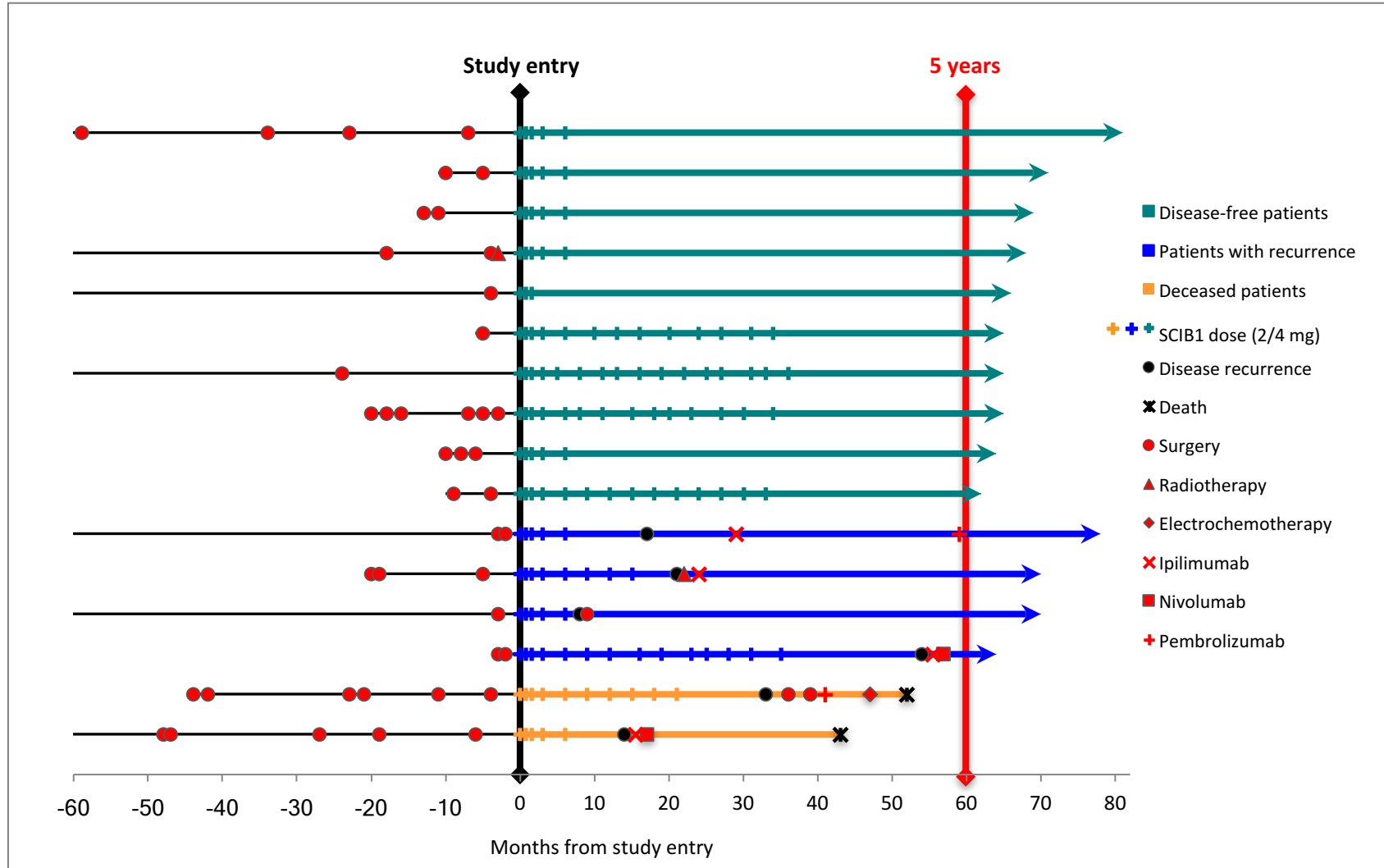
- ▶ Overall survival with SCIB1 treatment superior to historical survival rates
- ▶ 14 of 16 resected patients receiving 2-4 mg doses have survived for more than 5 years (February 2018)
- ▶ Melanoma recurrence rates are lower in SCIB1-treated patients than historical controls





SCIB1 IN MELANOMA PATIENTS

PATIENTS WITHOUT TUMOUR PRESENT AT STUDY ENTRY

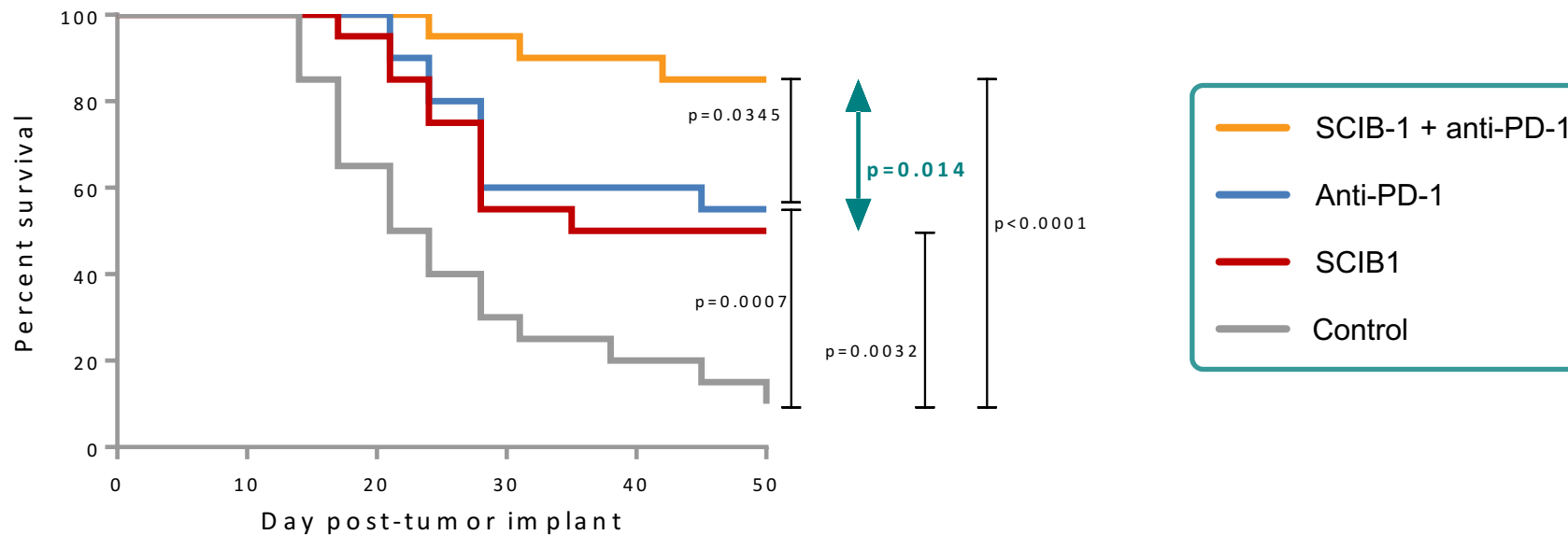




SCIB1 BOOSTS IMMUNE CHECKPOINT THERAPY

IN A MOUSE MELANOMA MODEL, SURVIVAL RATES WERE SIGNIFICANTLY BOOSTED WHEN ANTI-PD-1 THERAPY WAS COMBINED WITH SCIB1 TREATMENT

- ▶ Survival rates for SCIB1 ImmunoBody[®] monotherapy \approx anti-PD-1
- ▶ Monotherapy viable option for resected melanoma patients
- ▶ Combination therapy resulted in an 85% survival rate
- ▶ SCIB1 also upregulates PD-L1 expression and memory response

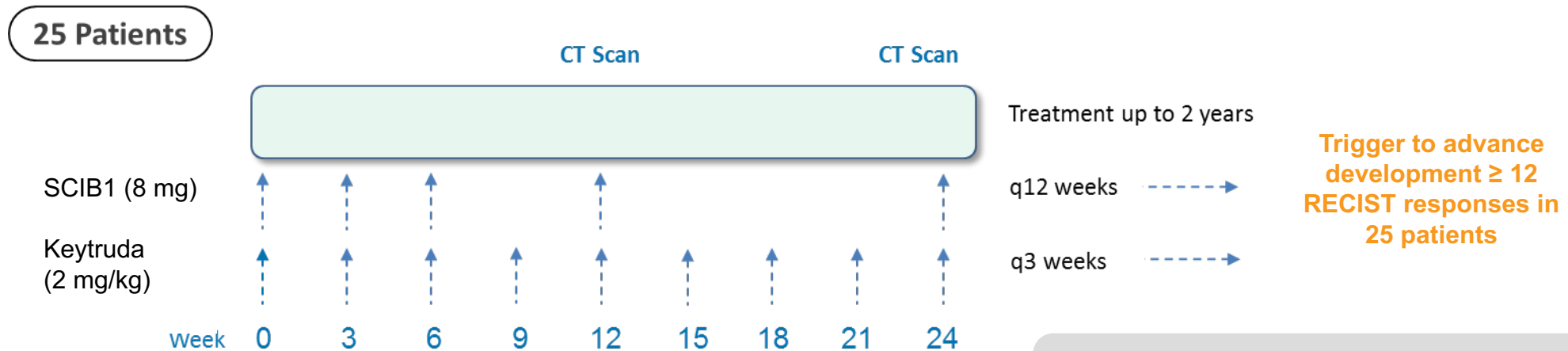




SCIB1 + CHECKPOINT INHIBITOR COMBINATION PHASE 2 TRIAL

PATIENT POPULATION

- ▶ Histologically confirmed, unresectable AJCC stage III or stage IV melanoma
- ▶ No prior systemic treatment for advanced disease
- ▶ Suitable for treatment with Keytruda (pembrolizumab), with measurable disease
- ▶ Part 1 safety run-in (n=6); Part 2 additional 19 patients; total = 25 patients



Assumptions

- ▶ Response rate to Keytruda = 30%
- ▶ Response rate of interest for combination = 55%



IMMUNOBODY®

SCIB1

- ▶ In combination with checkpoint inhibitors in patients with late stage disease to increase efficacy without compromising safety
- ▶ As monotherapy in patients with resected disease (adjuvant setting) to delay or prevent recurrence

SCIB2

- ▶ Lung cancer huge unmet medical need; deaths per year greater than melanoma, colon, breast and prostate cancers combined
- ▶ Checkpoint inhibitors less effective in lung cancer, with 80% of patients requiring a better SOC

MODITOPE®

Modi-1 & Modi-2

- ▶ siPTM vaccine class
- ▶ Innovative mechanism of action potentially targets all solid tumours
- ▶ Modi-1 and Modi-2 will target tumours that are unresponsive to existing immunotherapy (turning “cold” tumours to “hot”)
- ▶ Identification of Modi-specific TCRs provides a novel pathway for CD4-based TCR therapy



Operational

- ▶ SCIB2 clinical development partnership with Cancer Research UK (CRUK)-*December 2017*
- ▶ Moditope research Collaboration with BioNTech to develop T-cell receptor therapies- *January 2018*
- ▶ Licensing of ISA Pharmaceuticals Amplivant adjuvant technology for Modi-1- *February 2018*
- ▶ Contracting PolyPeptide Group for good manufacturing practice (GMP) manufacture of Modi-1- *March 2018*
- ▶ In-licensing of novel monoclonal antibodies against tumour associated glycans and antibody engineering technology- *April 2018*
- ▶ *Exercise of commercial option to Ichor TriGrid 2.0 electroporation delivery device for SCIB1*

Financial

- ▶ Cash at bank on 30th April 2018 was £10,303,168 (2017: £2,672.335)
- ▶ Overall loss for the year £4,194,509 (2017: £3,544,979)
- ▶ Share capital placing:
 - ▶ Placement £4.7m net of costs-May 2017
 - ▶ Placement £6.9m net of costs-April 2018
 - ▶ *Open Offer £1.1m net of costs-May 2018*
- ▶ *Exercise of share options by Ichor £143,307- July 2018*



IMMUNOBODY®

SCIB1

- ▶ SCIB1/checkpoint inhibitor Phase 2 US/UK combination study in late stage melanoma, planned to start 1H19, subject to regulatory submissions
 - ▶ Opening of IND
 - ▶ Commencement of the Phase 2 combination trial utilising Ichor TriGrid v2.0 electroporation device

SCIB2

- ▶ CRUK development activities for initiation of SCIB2 Phase 1/2 study for NSCLC

MODITOPE®

Modi-1

- ▶ Preparation for the First-In-Human study with Modi-1 in patients with TNBC, ovarian cancer and sarcoma planned to start CY19
- ▶ Identification of Modi-specific TCRs in collaboration with BioNTech

Modi-2

- ▶ Pre-clinical development for multiple solid tumour indications
- ▶ Extension of patent portfolio



FUTURE VALUE DRIVERS

2 PLATFORMS, 4 LEAD PRODUCTS + 5 CORE ACTIVITIES

CLINICAL DATA

- Generate meaningful clinical data to address unmet needs: clinical read-outs (SCIB1 Phase 2 & Modi-1 Phase 1/2 pt 1) anticipated in next 2 years

PIPELINE EXPANSION

- Extend utility of Moditope platform beyond Modi-1 and Modi-2 in association with key industry players e.g., TCRs
- Lead generation and optimisation of anti-glycan mAbs

TECHNOLOGY PARTNERSHIPS

- Evaluate and implement enabling technologies e.g., nano-vesicle delivery (Immunobody), and adjuvant (Moditope), to aid and de-risk development

CLINICAL PARTNERSHIPS

- Establish relationships with key opinion leaders and clinical networks to ensure utility in clinical practice e.g., CRUK and patient advocacy groups (Addario)

INDUSTRY PARTNERSHIPS

- Explore synergies with large Pharma/Biotech companies in identifying combination therapies for optimal outcomes e.g., checkpoint inhibitors





Contact

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